SEARCH REQUEST FORM (STIC)

	Requestor's Name: D	Pavid Lukton	Examiner numb		Date: 11 - 0 4				
	Art Unit: 1653	Phone number:		<u>Serial Nu</u> 9-581					
	Mail Box: 3-C-70	Examiner Rm: 3-		sults format:					
		****	*****						
	Title: Neurop	protective A	gents						
	Applicants: ST BRADLEY, MARK;	UNDSTROM, LA PRINGLE, AS		NOTTI, F	AUSTO;				
	Earliest Priority Date	: 12/16/97							
		* * * *	*****		W 11 20				
	Applicants are claimi	ng the compounds	s on the attached	sheet.	200 200 200				
	$R^2 = hydrogen or$	r alkyl or acyl or	R-NHCO-	(R = alkyl	or aryl);				
	R^3 = hydrogen or	r alkyl or acyl or	R-NHCO-	(R = alkyl	or aryl);				
	p = an integer of	3 or 4;							
	q = an integer of	3 or 4							

STAFF US		**************************************		************** rs and cost whe					
Scarcher:		NA Sequence (#)	STN						
Searcher Phone	#:	AA Sequence (#)	Dialog						
Searcher Locati	on:	Structure (#)	Questel/Orbit						
Date Searcher P	Date Searcher Pičked Up		Dr.Link	Dr.Link					
- zte Completeo	d:	Litigation	I_cxis/Nexis						
Searcher Prep &	Review Time:	Fulltext	Sequence Systems						
Clement Prop Ti	ime	Patent Family	www/Internet	olenen n nen Ine	8				
Online Libra	e same e e e e e e e e e e e e e e e e e e	Other							

PICLISPORTOR

Inventor Search

Lukton 09/581,397

11/05/2004

=> d ibib abs ind hitstr 142 1-3

L42 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

2003:753643 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 140:280736

Characterisation of a novel class of polyamine-based TITLE:

neuroprotective compounds

AUTHOR(S): Pringle, Ashley K.; Morrison, Barclay;

Bradley, Mark; Iannotti, Fausto;

Sundstrom, Lars E.

Clinical Neurosciences, University of Southampton, CORPORATE SOURCE:

Southampton, SO16 7PX, UK

Naunyn-Schmiedeberg's Archives of Pharmacology (2003), SOURCE:

368(3), 216-224

CODEN: NSAPCC; ISSN: 0028-1298

PUBLISHER: Springer-Verlag

DOCUMENT TYPE: Journal LANGUAGE: English Prolonged cerebral ischemia initiates complex intra- and inter-cellular

signalling cascades ultimately resulting in neuronal death. Well-characterised mediators of ischemic cell death are glutamate, free radicals and nitric oxide. Many drugs that block these mechanisms are neuroprotective in vitro, but have unfavorable side-effect profiles in man. We have recently demonstrated that the compound L-arginyl-3,4spermidine (L-Arg3,4) is neuroprotective in vitro through an interaction with several of these mechanisms, and prevents ischemic neurodegeneration

in vivo with no gross side effects. In this study, we have used solid-phase combinatorial chemical, to synthesize a number of analogs of L-Arg3,4, and investigate the structure-activity relationship using an in vitro, organotypic hippocampal slice culture model of cerebral ischemia. A number of mol. features were identified which were essential for the neuroprotective activity including the requirement for a pos. charge and an amino acid in the L-configuration. Relatively minor alterations to both the terminal arginine and polyamine moieties significantly attenuated neuroprotective efficacy. Our data implies that these compds. are neuroprotective through a currently undefined mechanism rather than non-specific ionic interactions described previously for other polyamine-containing compds.

CC1-3 (Pharmacology)

structure activity neuroprotectant polyamine ischemia brain hippocampus ST

TΤ Brain

(hippocampus; structure and neuroprotective activity of polyamine-based L-arginyl-3,4-spermidine analogs)

TΤ Brain, disease

(ischemia; structure and neuroprotective activity of polyamine-based L-arginyl-3,4-spermidine analogs)

ΙT Cytoprotective agents

(neuroprotective; structure and neuroprotective activity of polyamine-based L-arginyl-3,4-spermidine analogs)

ITStructure-activity relationship

(structure and neuroprotective activity of polyamine-based L-arginyl-3,4-spermidine analogs)

IT 134950-93-9 134951-15-8 141997-14-0

191277-14-2 227758-27-2 227758-28-3

227758-29-4 227758-36-3 227758-40-9

227758-41-0 675606-34-5 675606-35-6

675606-36-7 675606-37-8 675606-38-9

675606-39-0 675606-40-3

RL: PAC (Pharmacological activity); BIOL (Biological study) (structure and neuroprotective activity of polyamine-based

L-arginyl-3,4-spermidine analogs) 134950-93-9 134951-15-8 141997-14-0 IT 191277-14-2 227758-27-2 227758-28-3 227758-29-4 227758-36-3 227758-40-9 227758-41-0 675606-34-5 675606-35-6 675606-36-7 675606-37-8 675606-38-9 675606-39-0 675606-40-3 RL: PAC (Pharmacological activity); BIOL (Biological study) (structure and neuroprotective activity of polyamine-based L-arginyl-3,4-spermidine analogs) 134950-93-9 HCAPLUS RN Pentanamide, 2-amino-5-[(aminoiminomethyl)amino]-N-[4-[(3-CN aminopropyl)amino]butyl]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 $(CH_2)_3$
 N
 H
 $(CH_2)_4$
 N
 H
 $(CH_2)_4$
 N
 H
 NH_2

RN 134951-15-8 HCAPLUS
CN Hexanamide, 2,6-diamino-N-[3-[(4-aminobutyl)amino]propyl]-, (2S)- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 $(CH_2)_4$
 N
 H
 $(CH_2)_3$
 N
 H
 $(CH_2)_4$
 NH_2

RN 141997-14-0 HCAPLUS
CN Pentanamide, 2-amino-5-[(aminoiminomethyl)amino]-N-[3-[(3-aminopropyl)amino]propyl]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 N
 H
 $(CH_2)_3$
 N
 H
 $(CH_2)_3$
 N
 H
 $(CH_2)_3$
 N
 H
 N
 N
 H

RN 191277-14-2 HCAPLUS
CN Pentanamide, 2-amino-N-[3-[(4-aminobutyl)amino]propyl]-5[(aminoiminomethyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

$$_{\text{H}_{2}\text{N}}$$
 (CH₂) 4 (CH₂) 3 N (CH₂) 3 N NH₂

RN 227758-27-2 HCAPLUS
CN Hexanamide, 2,6-diamino-N-[3-[(4-aminobutyl)amino]propyl]-, (2R)- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.

$$(CH_2)_4$$
 $(CH_2)_3$ $(CH_2)_4$ $(CH_2)_4$

RN 227758-28-3 HCAPLUS
CN Pentanamide, 2,5-diamino-N-[3-[(4-aminobutyl)amino]propyl]-, (2S)- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 $(CH_2)_3$
 N
 H
 $(CH_2)_4$
 NH_2

RN 227758-29-4 HCAPLUS
CN Pentanamide, 2,5-diamino-N-[3-[(4-aminobutyl)amino]propyl]-, (2R)- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.

$$(CH_2)_3$$
 $(CH_2)_3$ $(CH_2)_4$ $(CH_2)_4$

RN 227758-36-3 HCAPLUS
CN Pentanamide, 2-amino-N-[3-[(4-aminobutyl)amino]propyl]-5[(aminocarbonyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

$$H_2N$$
 $(CH_2)_4$
 N
 H
 $(CH_2)_3$
 N
 H
 $(CH_2)_3$
 N
 H
 NH_2

RN 227758-40-9 HCAPLUS

CN Pentanediamide, 2-amino-N1-[3-[(4-aminobutyl)amino]propyl]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 N
 H_2
 N
 H_2
 N
 H
 H
 N
 H
 N
 H
 N
 H
 N
 H
 N
 H

RN 227758-41-0 HCAPLUS

CN Pentanamide, 2-amino-N-[4-[(4-aminobutyl)amino]butyl]-5-[(aminoiminomethyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$(CH_2)_4$$
 $(CH_2)_4$
 $(CH_2)_4$

RN 675606-34-5 HCAPLUS

CN 1H-Imidazole-4-propanamide, α -amino-N-[3-[(4-aminobutyl)amino]propyl]-, (α S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 675606-35-6 HCAPLUS

CN Hexanamide, 2,6-diamino-N-[4-[(4-aminobutyl)amino]butyl]-, (2S)- (9CI) (CA INDEX NAME)

RN 675606-36-7 HCAPLUS

CN Pentanamide, 2-amino-5-[(aminoiminomethyl)amino]-N-(8-aminooctyl)-, (2S)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 675606-37-8 HCAPLUS

CN Pentanamide, N-[3-[(4-aminobutyl)amino]propyl]-5-[(aminoiminomethyl)amino]-2-[(phenylmethyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 H_1
 H_2N
 H_1
 H_2N
 H_1
 H_2
 H_3
 H_4
 H_4
 H_5
 H_5
 H_6
 H_7
 H

RN 675606-38-9 HCAPLUS

CN Carbamic acid, [(1S)-1-[[[3-[(4-aminobutyl)amino]propyl]amino]carbonyl]-4-[(aminoiminomethyl)amino]butyl]-, phenylmethyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 H_1
 H_2N
 H_2N
 H_2N
 H_1
 H_2N
 H_2N
 H_2N
 H_1
 H_2N
 H_2N
 H_1
 H_2N
 H_2N
 H_1
 H_2N
 H_2N
 H_1
 H_2N
 H_2N
 H_2N
 H_1
 H_1
 H_1
 H_2N
 H_1
 H_1

RN 675606-39-0 HCAPLUS

CN Pentanamide, N-[3-[(4-aminobutyl)amino]propyl]-5-[(aminoiminomethyl)amino]-2-[(phenylsulfonyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 N_H
 $(CH_2)_3$
 N_H
 $(CH_2)_4$
 N_H
 N_H

675606-40-3 HCAPLUS RN

Pentanamide, 2-(acetylamino)-N-[3-[(4-aminobutyl)amino]propyl]-5-CN [(aminoiminomethyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_{2}N$$
 $H_{2}N$
 $H_{3}N$
 $H_{4}N$
 $H_{5}N$
 H

REFERENCE COUNT:

41 THERE ARE 41 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L42 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2003:4310 HCAPLUS

DOCUMENT NUMBER:

139:30604

TITLE:

L-Arginyl-3,4-spermidine is neuroprotective in several.

in vitro models of neurodegeneration and in vivo ischaemia without suppressing synaptic transmission

AUTHOR(S):

Morrison, Barclay, III; Pringle, Ashley K.; McManus, Terence; Ellard, John; Bradley, Mark ; Signorelli, Francesco; Iannotti, Fausto;

Sundstrom, Lars E.

CORPORATE SOURCE:

Division of Clinical Neurosciences, School of

Medicine, Bassett Crescent East, University of Southampton, Southampton, SO16 7PX, UK

British Journal of Pharmacology (2002), 137(8),

1255-1268

CODEN: BJPCBM; ISSN: 0007-1188

PUBLISHER:

SOURCE:

Nature Publishing Group

DOCUMENT TYPE:

Journal

LANGUAGE:

English

1 Stroke is the third most common cause of death in the world, and there is a clear need to develop new therapeutics for the stroke victim. To address this need, we generated a combinatorial library of polyamine compds. based on sFTX-3.3 toxin from which L-Arginyl-3,4-spermidine (L-Arg-3,4) emerged as a lead neuroprotective compound In the present study, we have extended earlier results to examine the compound's neuroprotective actions in greater detail. 2 In an in vitro ischemia model, L-Arg-3,4 significantly reduced CA1 cell death when administered prior to induction of 60 min of ischemia as well as when administered immediately after ischemia. Surprisingly, L-Arg-3,4 continued to prevent cell death significantly when administration was delayed for as long as 60 min after ischemia. 3 L-Arg-3,4 significantly reduced cell death in excitotoxicity models mediated by glutamate, NMDA, AMPA, or kainate. Unlike glutamate receptor antagonists, 300 µM L-Arg-3,4 did not suppress synaptic transmission as measured by evoked responses in acute hippocampal slices. 4 L-Arg-3,4 provided significant protection, in vitro, in a superoxide mediated injury model and prevented an increase of superoxide production after AMPA or NMDA stimulation. It also decreased nitric oxide production after in vitro ischemia and NMDA stimulation, but did so without inhibiting nitric oxide synthase directly. 5 Furthermore, L-Arg-3,4 was significantly neuroprotective in an in vivo model of global forebrain ischemia, without any apparent neurol. side-effects. 6 Taken together, these results demonstrate that L-Arg-3,4 is protective in several models of neurodegeneration and may have potential as a new therapeutic compound for the treatment of stroke, trauma, and other neurodegenerative diseases.

CC 1-11 (Pharmacology)

ST arginylspermidine neuroprotective forebrain ischemia stroke

IT Glutamate receptors

RL: BSU (Biological study, unclassified); BIOL (Biological study) (AMPA-binding; arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT Brain, disease

(forebrain, ischemia; arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT Brain

(hippocampus, sector CA1, cell death inhibition; arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT Cytoprotective agents

(neuroprotective; arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT Toxicity

(neurotoxicity, excitotoxicity; arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT Brain, disease

(stroke; arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT Neurotransmission

(synaptic; arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT Nerve

(toxicity, excitotoxicity; arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT 56-86-0, L-Glutamic acid, biological studies 487-79-6,

Kainic acid 6384-92-5 10102-43-9, Nitric oxide,

biological studies 11062-77-4, Superoxide

RL: ADV (Adverse effect, including toxicity); BIOL (Biological study) (arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT 191277-14-2

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

IT 56-86-0, L-Glutamic acid, biological studies 487-79-6,

Kainic acid 6384-92-5 10102-43-9, Nitric oxide,

biological studies 11062-77-4, Superoxide

RL: ADV (Adverse effect, including toxicity); BIOL (Biological study) (arginylspermidine is neuroprotective in several in vitro models of neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

RN 56-86-0 HCAPLUS

CN L-Glutamic acid (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 487-79-6 HCAPLUS

CN 3-Pyrrolidineacetic acid, 2-carboxy-4-(1-methylethenyl)-, (2S,3S,4S)-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

$$HO_2C$$
 S
 S
 S
 S
 S
 S
 S
 S

RN 6384-92-5 HCAPLUS

CN D-Aspartic acid, N-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 10102-43-9 HCAPLUS

CN Nitrogen oxide (NO) (8CI, 9CI) (CA INDEX NAME)

N = 0

RN 11062-77-4 HCAPLUS

CN Superoxide (8CI, 9CI) (CA INDEX NAME)

o = o

IT 191277-14-2

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (arginylspermidine is neuroprotective in several in vitro models of

neurodegeneration and in vivo ischemia without suppressing synaptic transmission)

RN 191277-14-2 HCAPLUS

CN Pentanamide, 2-amino-N-[3-[(4-aminobutyl)amino]propyl]-5-[(aminoiminomethyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 $(CH_2)_4$
 N
 H
 $(CH_2)_3$
 N
 H
 NH_2
 NH_2

REFERENCE COUNT:

THERE ARE 45 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L42 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

1999:404918 HCAPLUS

DOCUMENT NUMBER:

131:59135

TITLE:

Preparation of amino acid derivatives as

neuroprotective agents

INVENTOR(S):

Pringle, Ashley Ker; Bradley, Mark
; Sundstrom, Lars Eric; Iannotti,

Fausto

PATENT ASSIGNEE(S):

University of Southampton, UK

SOURCE:

PCT Int. Appl., 53 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

1

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.		KI	ND	DATE			A	PPLI	CATIO	N NC	Э.	DATE					
WO 9931049			Α	A1 19990624				WO 1998-GB3775				5	19981216				
	W:	AL,	AM,	AT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,	DE,
		DK,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,	HU,	ID,	ΙL,	IN,	IS,	JP,
				•										MD,			
														SK,			
		TR,	TT,	UA,	UG,	US,	UZ,	VN,	YU,	ZW,	ΑM,	ΑZ,	BY,	KG,	KΖ,	MD,	RU,
		ТJ,	TM														
	RW:	GH,	GM,	KE,	LS,	MW,	SD,	SZ,	UG,	ZW,	ΑT,	BE,	CH,	CY,	DE,	DK,	ES,
		FΙ,	FR,	GB,	GR,	ΙE,	ΙT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,
		CM,	•	GN,				•	•	•							
CA 2315258		AA 19990624				CA 1998-2315258				58	19981216						
AU 9915717		Α	1	19990705			AU 1999-15717				19981216						
ΑU	AU 739296		В	2	2001	1011											
EP 1040096		Α	1	20001004			EP 1998-960031			1	19981216						
EΡ	1040	096		В	1	2003	0709										

```
AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
     JP 2002508349
                       T2
                            20020319
                                           JP 2000-538979
                                                             19981216
                                           NZ 1998-505110
                            20030131
     NZ 505110
                       Α
                                                             19981216
                       C2
                                           RU 2000-116112
                            20030527
                                                             19981216
     RU 2205177
                                           AT 1998-960031
                            20030715
                                                             19981216
     AT 244698
                       E
                                           PT 1998-960031
                       т
                            20030930
                                                             19981216
     PT 1040096
                                           ES 1998-960031
     ES 2201563
                       Т3
                            20040316
                                                             19981216
                            20000622
                                           CA 1999-2355880
                                                             19990616
     CA 2355880
                       AΑ
                                           WO 1999-GB1719
     WO 2000035941
                       Α2
                            20000622
                                                             19990616
                            20011004
     WO 2000035941
                       AЗ
         W: CA, US
         RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE
                                            EP 1999-936759
                                                             19990616
                            20011017
     EP 1144434
                       A2
                            20020529
     EP 1144434
                       ΑЗ
                     CH, DE, DK, ES, FR, GB, IT, LI, NL, SE, IE
         R: AT, BE,
                                           NO 2000-3075
                                                             20000615
                            20000815
     NO 2000003075
                       Α
                            20031003
                                            HK 2000-108125
                                                             20001215
     HK 1029331
                       Α1
                                        GB 1997-26569
                                                             19971216
PRIORITY APPLN. INFO .:
                                                          Α
                                        WO 1998-GB3775
                                                             19981216
                                                          W
                                        WO 1999-GB1719
                                                             19990616
                                                          W
                         MARPAT 131:59135
OTHER SOURCE(S):
    Amino acid derivs. Q-Ra-C*H(NR2R3)CO-Zn-NR1-Rb-NH-Rc-NH-W [Q = amidino,
     cyano, or amino group; Ra, Rb, Rc = (un) substituted alkylene, alkenylene;
     R2, R3 = H, R, RCO, RO2C, RNHCO (R = (un) substituted alkyl or aryl); the
     chiral atom indicated by the asterisk is in the L configuration; Z is an
     amino acid residue; n = 0, 1; R1 = H, (un) substituted alkyl or aryl; W =
     H, alkyl, aryl] were prepared as neuroprotectants.
                                                         Thus,
     N1-L-arginylspermidine, prepared by coupling of resin-bound spermidine
     derivative with protected arginine, followed by deprotection/cleavage using
     TFA-phenol-water-triisopropylsilane-1,2-ethanedithiol, showed 99.4 %
     protection (relative to control hypoxia in CA1 pyramidal cell layer).
IC
     ICM C07C237-10
     ICS C07C257-14; A61K031-155; A61K031-16
CC
     34-2 (Amino Acids, Peptides, and Proteins)
     arginylspermidine prepn neuroprotectant; spermidine arginyl prepn
ST
     neuroprotectant
IT
     Structure-activity relationship
        (neuroprotectant; preparation of amino acid derivs. as neuroprotective
        agents)
     Cytoprotective agents
IT
        (neuroprotectants; preparation of amino acid derivs. as neuroprotective
        agents)
IT
     Ischemia
        (preparation of amino acid derivs. as neuroprotective agents)
TT
     Amino acids, preparation
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of amino acid derivs. as neuroprotective agents)
IT
     134951-15-8P 191277-14-2P 191277-15-3P
     227758-27-2P 227758-28-3P 227758-29-4P
     227758-31-8P 227758-32-9P 227758-33-0P
     227758-34-1P 227758-35-2P 227758-36-3P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of amino acid derivs. as neuroprotective agents)
IT
     227758-40-9 227758-41-0 227767-50-2
```

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(preparation of amino acid derivs. as neuroprotective agents)

IT 110-60-1, 1,4-Butanediamine 156-87-6

227758-37-4D, resin-bound

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of amino acid derivs. as neuroprotective agents)

IT 227758-39-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of amino acid derivs. as neuroprotective agents)

IT 134951-15-8P 191277-14-2P 191277-15-3P

227758-27-2P 227758-28-3P 227758-29-4P

227758-31-8P 227758-32-9P 227758-33-0P

227758-34-1P 227758-35-2P 227758-36-3P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of amino acid derivs. as neuroprotective agents)

RN 134951-15-8 HCAPLUS

CN Hexanamide, 2,6-diamino-N-[3-[(4-aminobutyl)amino]propyl]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 $(CH_2)_4$
 N
 H
 $(CH_2)_3$
 N
 H
 $(CH_2)_4$
 NH_2

RN 191277-14-2 HCAPLUS

CN Pentanamide, 2-amino-N-[3-[(4-aminobutyl)amino]propyl]-5-[(aminoiminomethyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$(CH_2)_4$$
 $(CH_2)_3$
 $(CH_2)_3$
 $(CH_2)_3$
 $(CH_2)_3$
 $(CH_2)_4$
 $(CH_2)_4$

RN 191277-15-3 HCAPLUS

CN Pentanamide, 2-amino-N-[3-[(4-aminobutyl)amino]propyl]-5-[(aminoiminomethyl)amino]-, (2R)- (9CI) (CA INDEX NAME)

$$H_2N$$
 $(CH_2)_4$
 N
 H
 $(CH_2)_3$
 N
 H
 NH_2
 NH_2

RN 227758-27-2 HCAPLUS

CN Hexanamide, 2,6-diamino-N-[3-[(4-aminobutyl)amino]propyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 $(CH_2)_4$
 R
 N
 H
 $(CH_2)_3$
 N
 H
 $(CH_2)_4$
 NH_2

RN 227758-28-3 HCAPLUS

CN Pentanamide, 2,5-diamino-N-[3-[(4-aminobutyl)amino]propyl]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 $(CH_2)_3$
 N
 H
 $(CH_2)_4$
 NH_2

RN 227758-29-4 HCAPLUS

CN Pentanamide, 2,5-diamino-N-[3-[(4-aminobutyl)amino]propyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$(CH_2)_3$$
 $(CH_2)_4$ $(CH_2)_4$

RN 227758-31-8 HCAPLUS

CN L-Phenylalaninamide, L-arginyl-N-[3-[(4-aminobutyl)amino]propyl]-,
 tetrakis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 227758-30-7

CMF C22 H40 N8 O2

Absolute stereochemistry.

$$H_2N$$
 NH
 $(CH_2)_3$
 S
 NH_2
 $(CH_2)_3$
 $(CH_2)_4$
 $(CH_2)_4$
 $(CH_2)_4$
 $(CH_2)_4$
 $(CH_2)_4$

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 227758-32-9 HCAPLUS

CN L-Phenylalaninamide, N6-(aminoiminomethyl)-L-lysyl-N-[3-[(4-aminobutyl)amino]propyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 NH
 $(CH_2)_4$
 S
 NH_2
 O
 $(CH_2)_3$
 NH_2
 O
 $(CH_2)_4$
 NH_2

RN 227758-33-0 HCAPLUS

CN L-Tyrosinamide, L-arginyl-N-[3-[(4-aminobutyl)amino]propyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

HO
$$\begin{pmatrix} CH_2 \end{pmatrix} 3 \begin{pmatrix} CH_2 \end{pmatrix} 4 \end{pmatrix}$$
 $\begin{pmatrix} CH_2 \end{pmatrix} 4 \end{pmatrix}$
 $\begin{pmatrix} CH_2 \end{pmatrix} 3 \begin{pmatrix} CH_2 \end{pmatrix} 4 \end{pmatrix}$
 $\begin{pmatrix} CH_2 \end{pmatrix} 3 \begin{pmatrix} CH_2 \end{pmatrix} 3 \begin{pmatrix} CH_2 \end{pmatrix} 3 \end{pmatrix}$
 $\begin{pmatrix} CH_2 \end{pmatrix} 3 \begin{pmatrix} CH_2 \end{pmatrix} 3 \begin{pmatrix}$

RN 227758-34-1 HCAPLUS

CN L-Tryptophanamide, L-arginyl-N-[3-[(4-aminobutyl)amino]propyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 227758-35-2 HCAPLUS

CN Glycinamide, L-arginyl-N-[3-[(4-aminobutyl)amino]propyl]-2-phenyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 227758-36-3 HCAPLUS

CN Pentanamide, 2-amino-N-[3-[(4-aminobutyl)amino]propyl]-5-[(aminocarbonyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

IT 227758-40-9 227758-41-0 227767-50-2

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(preparation of amino acid derivs. as neuroprotective agents)

RN 227758-40-9 HCAPLUS

CN Pentanediamide, 2-amino-N1-[3-[(4-aminobutyl)amino]propyl]-, (2S)- (9CI) (CA INDEX NAME)

$$H_2N$$
 N
 H_2
 N
 H_2
 N
 H
 N
 H
 N
 H
 N
 H
 N
 H
 N
 H

RN 227758-41-0 HCAPLUS

CN Pentanamide, 2-amino-N-[4-[(4-aminobutyl)amino]butyl]-5-[(aminoiminomethyl)amino]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$H_2N$$
 $(CH_2)_4$
 N
 H
 $(CH_2)_4$
 N
 H
 $(CH_2)_4$
 N
 H
 NH_2

RN 227767-50-2 HCAPLUS

CN Pyridinepropanamide, α -amino-N-[3-[(4-aminobutyl)amino]propyl]-(9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & \text{O} & \text{NH}_2 \\ & \parallel & \parallel \\ & \text{H}_2\text{N--} \text{(CH}_2)}_4 - \text{NH--} \text{(CH}_2)}_3 - \text{NH--} \text{C--} \text{CH--} \text{CH}_2 - \text{D1} \end{array}$$

IT 110-60-1, 1,4-Butanediamine 156-87-6

227758-37-4D, resin-bound

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of amino acid derivs. as neuroprotective agents)

RN 110-60-1 HCAPLUS

CN 1,4-Butanediamine (8CI, 9CI) (CA INDEX NAME)

 $H_2N - (CH_2)_4 - NH_2$

RN 156-87-6 HCAPLUS

CN 1-Propanol, 3-amino- (8CI, 9CI) (CA INDEX NAME)

H2N-CH2-CH2-CH2-OH

RN 227758-37-4 HCAPLUS

CN 13-0xa-2,6,11-triazapentadecanoic acid, 6-[2-amino-1-[4-[(benzoyloxy)carbonyl]phenoxy]-2-oxoethyl]-14,14-dimethyl-12-oxo-,

9H-fluoren-9-ylmethyl ester (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

IT 227758-39-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of amino acid derivs. as neuroprotective agents)

RN 227758-39-6 HCAPLUS

CN 1,3-Cyclohexanedione, 2-[1-[(3-hydroxypropyl)amino]ethylidene]-5,5-dimethyl- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=>